

Inventor: Kathryn Lindsay BALL et al.

Title: METHODS AND MEANS FOR INHIBITION OF CDK4

ACTIVITY

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Peptide 1 M S E P A G D V R Q N P C G S K A Peptide 2. V D S Ε \mathbf{L} S Q R D Peptide 3. S D Ι Ε Α R G С Peptide 4. R \mathbf{T} E Т Ρ L Ε G E D Peptide 5. G G L K L Peptide 6. R Ε G G D R Peptide 7. G G R R Ρ Α Q G \mathbf{T} Α Peptide 8. Α E EL S L S C \mathbf{T} ₽ R S D V Peptide 9. G S P R S G Ε Α Ε Ρ G G Ρ G D S G Peptide 10. KRR Т S Μ \mathbf{T} D F Y Η S K R R F S L Ι Peptide 11. \mathbf{T} Y H S K RRLIFS KRKP Μ

Fig. 1A

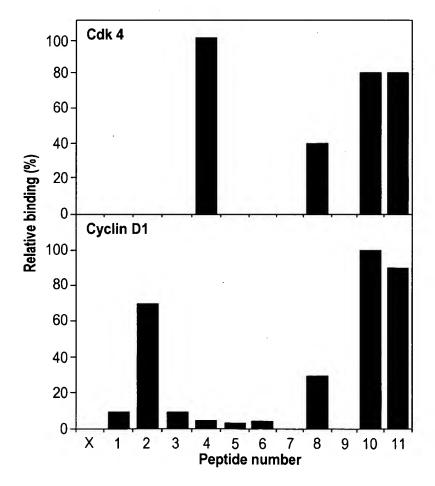


Fig. 1B

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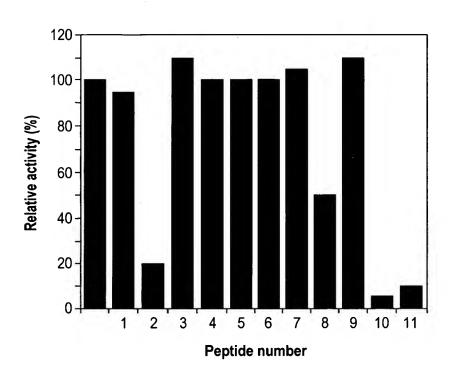
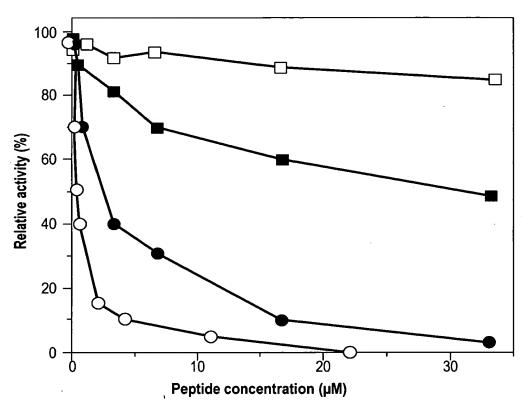


Fig. 2

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Peptide number	l _{0.5}
4 (□)	-
8 (■)	>34.0µM
2 (•)	2.0µM
10 (0)	0.1µM

Fig. 3

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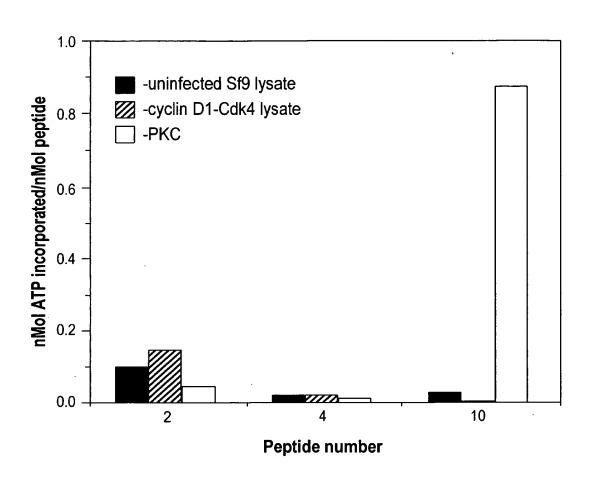


Fig. 4

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a -	PRSGEQAEGSPGGPGDSQGR
b -	EQAEGSPGGPGDSQGRKRRQ
C -	GSPGGPGDSQGRKRRQTSMT
d -	GPGDSQGRKRRQTSMTDFYH
e -	SQGRKRRQTSMTDFYHSKRR
peptide 10-	KRRQTSMTDFYHSKRRLIFS
f-	TSMT DFYHSKRRLIFS KRKF
g -	DFYHSKRRLIFSKRKF

Fig. 5

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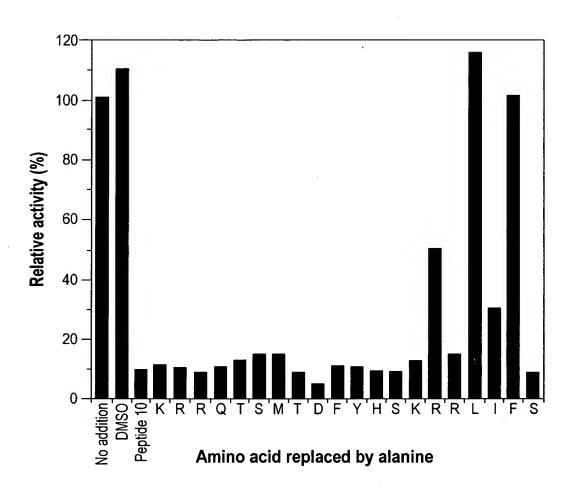
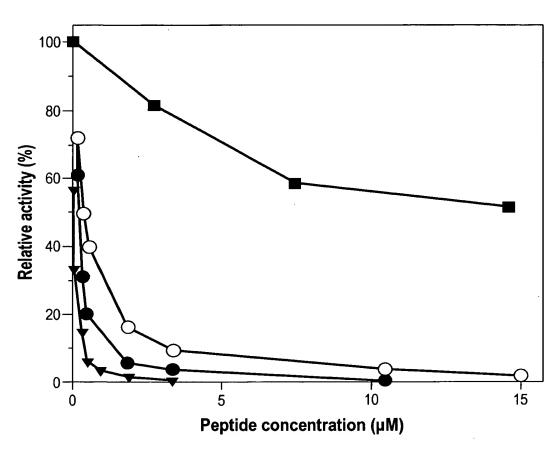


Fig. 6

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Peptide/protein	l _{0.5}
p16 ^{INK4} -peptide (■)	16.3 µM
10 (0)	0.1 µM
Asp→Ala mutant (●)	46 nM
Full-length p21 ^{WAF1} (▼)	11 nM

Fig. 7

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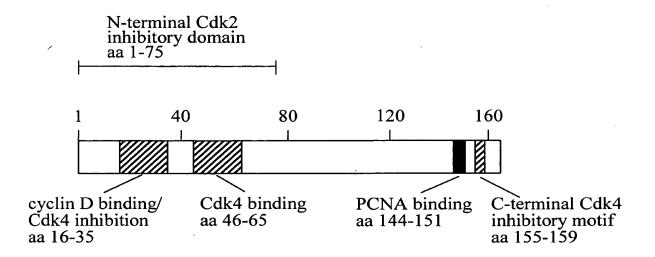


Fig. 8

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Peptide I KRRQTSATDFYHSKRRLIFSRQIKIWFQNRRMKWKK

Peptide II KRRLIFSKRQIKIWFQNRRMKWKK

Peptide III RQTSMTDFYHSKRRRQIKIWFQNRRMKWKK

Fig. 9A

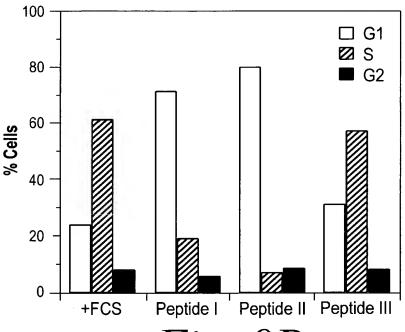


Fig. 9*B*

Phase Cell-cycle distribution (%) MRC5 MCF7 Control Peptide I Control Peptide I 37 42 68 G1 66 S 36 16 38 14 G2/M 16 13 10 12

Fig. 9C

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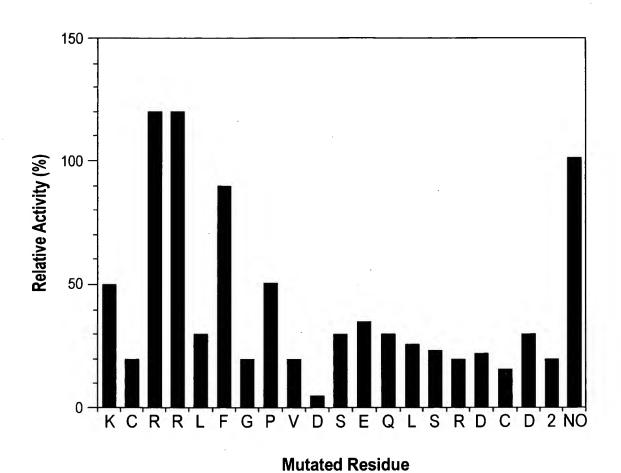


Fig. 10